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09/888,309

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## Applicati n Number TRANSMITTAL June 21, 2001 **Filing Date FORM** Melissa K. Carpenter, et al. **First Named Inventor** (to be used for all correspondence after initial filing) Group Art Unit 1632 Thaian N. Ton **Examiner Name** 090/002 9 Total Number of Pages in This Submission Attorney Docket Number **ENCLOSURES** (check all that apply) After Allowance Communication **Assignment Papers** Fee Transmittal Form (for an Application) to Group Appeal Communication to Board Fee Attached Drawing(s) of Appeals and Interferences Appeal Communication to Group Licensing-related Papers Amendment / Reply (Appeal Notice, Brief, Reply Brief) Petition After Final Proprietary Information Petition to Convert to a Affidavits/declaration(s) **Provisional Application** Status Letter Power of Attorney, Revocation Change of Correspondence Address (1 page) Other Enclosure(s) (please Extension of Time Request identify below): **Terminal Disclaimer** Preliminary Amendment (6 pages), Version Express Abandonment Request with Markings to Show Changes Made (2 Request for Refund pages) Information Disclosure Statement CD, Number of CD(s)\_ Certified Copy of Priority Document(s) Remarks Response to Missing Parts/ Incomplete Application Response to Missing Parts under 37 CFR 1.52 or 1.53 SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT Firm J. Michael Schiff, Registration No. 40,253 Individual name Signature Date -churc 15, 2002 CERTIFICATE OF HAND DELIVERY I hereby certify that this correspondence is being hand delivered to the Commissioner for Patents, Washington, DC 20231 on this date: February 19, 2002 Shari Hall White Typed or printed name February 19, 2002

Date

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I hereby certify that this paper is being delivered by hand to the U.S. Patent Office in accordance with 37 CFR § 1.6(b), addressed to the Assistant Commissioner for Patents, Washington, D.C. 20231, on the date indicated.

Name

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In the application of: M. Carpenter et al.

09/888,309 Serial No.:

Filing Date: June 21, 2001

For: DOPAMINERGIC NEURONS OBTAINED

FROM HUMAN EMBRYONIC STEM CELLS

Art Unit:

1632

Examiner:

Thaian N. Ton, Ph.L

PRELIMINARY AMENDMENT

**Assistant Commissioner for Patents** Washington, D.C. 20231

Dear Sir:

Applicants understand that this application has not yet been examined on the merits. Accordingly, please enter the enclosed amendments and remarks into the application, pursuant to 37 CFR § 1.115(b)(1).

PATENT 09/888,309 Docket: 090/002

## **AMENDMENTS**

Please delete the current TITLE of the application and replace it with the following:



## DOPAMINERGIC NEURONS OBTAINED FROM HUMAN EMBRYONIC STEM CELLS

Please cancel claims 1-20 without prejudice, and replace them with the following new claims:

23. A method for producing a neural cell population from human embryonic stem (hES) cells, comprising culturing progeny of the hES cells in a medium containing one or more added TGF-β Superfamily Antagonists so as to produce a population in which at least 50% of the cells express either polysialylated NCAM or β-tubulin III.



- 24. The method of claim 23, wherein the progeny are cultured in a medium containing noggin.
- 25. The method of claim 23, wherein the progeny are cultured in a medium containing follistatin.
- 26. The method of claim 23, wherein the medium further contains a neurotrophin.
- 27. The method of claim 26, wherein the neurotrophin is NT-3 or BDNF.
- 28. The method of claim 23, wherein the medium further contains a combination of factors selected from differentiation factors, neurotrophic factors, and survival factors.
- 29. The method of claim 23, comprising differentiating the hES cells by plating them onto a solid surface without forming embryoid bodies or cell aggregates.
- 30. The method of claim 29, wherein the solid surface comprises fibronectin or a polycation.
- 31. The method of claim 23, wherein at least 10% of the MAP-2 positive cells in the produced

PATENT 09/888,309 Docket: 090/002

population express tyrosine hydroxylase.

- 32. The method of claim 23, further comprising combining the cell population with a compound, determining any phenotypic or metabolic changes in the cell that result from contact with the compound, and correlating the change with cellular toxicity or modulation caused by the compound.
- 33. The method of claim 23, further comprising identifying an mRNA expressed at a different level in the neural cell population, relative to the level in undifferentiated hES cells; and preparing a polynucleotide comprising a nucleotide sequence of at least 30 consecutive nucleotides contained in the identified mRNA.
- 34. A set of two cultured cell populations, consisting of:

a first cell population comprising undifferentiated cells from a line of human embryonic stem (hES) cells; and

a second cell population, comprising progeny of the hES cells in a medium containing one or more added TGF-β Superfamily Antagonists.

- 35. A set of two isolated cell populations, consisting of:
  - a first cell population comprising undifferentiated cells from a line of human embryonic stem (hES) cells; and

a second cell population, comprising at least ~10% hES derived neural cells, identifiable by the criteria that they are progeny of said hES cell line and express both MAP-2 and tyrosine hydroxylase.

36. The set of cell populations of claim 35, wherein the second population has been produced from cells of the first population (or their progeny) by the method of claim 23.

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